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# Intravitreal Aflibercept Exposure During Early Pregnancy: A Case-Based Risk Evaluation

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### Introduction:

- Aflibercept is a VEGF inhibitor used for retinal diseases (1).
  - VEGF is essential in fetal vascular development.
- Poorly information about risk of aflibercept during early pregnancy (1st trimester).

VEGF in Adults	VEGF in Fetal Development
Promotes abnormal blood vessels (eye diseases)	Essential for placental and organ formation



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# **Methods**

- Retrospective study at Marmara
   University Teratology Information
   Service.
- Detailed histories, follow-ups, fetal and neonatal data collected.



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# Results Table of Aflibercept Use During Pregnancy

Parameter	Case 1	Case 2
Age	44 years old	34 years old
Indication	Diabetic retinopathy	Choroidal neovascularization
<b>Aflibercept Injection Timing</b>	Weeks 7, 10, and 17	Week 8
Comorbidities	Malignant hypertension	None reported
Medications	Perindopril, Amlodipine, etc.	None specified
Smoking Status	Heavy smoker (30–40 cigs/day until week 17)	Non-smoker
Delivery	Cesarean section at 37 weeks	Vaginal delivery at 38 weeks
Infant Outcome	Healthy baby; no malformations; normal growth (50–90th percentiles)	Healthy infant; normal birth weight and growth percentiles

# Conclusion

- Animal studies show fetal toxicity, but human exposure is low after intravitreal use (2).
  - No adverse outcomes were seen in these two cases.
- The first case involved complicating factors like hypertension and drug interactions.

# **Implications**

- Aflibercept may have lower teratogenic risk than expected (3-5).
  - More clinical data are needed to confirm safety.
- Useful for teratology risk assessment and patient counseling.

#### References

- 1. Wells, J. A., Glassman, A. R., Ayala, A. R., Jampol, L. M., Aiello, L. P., Antoszyk, A. N., ... & Beck, R. W. (2015). Aflibercept, bevacizumab, or ranibizumab for diabetic macular edema. The New England journal of medicine, 372(13), 1193-1203.
  - 2. Regeneron Pharmaceuticals. 2024. Eylea labeling. https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=f96cfd69-da34-41ee-90a9-610a4655cd1c&type=display
    3. Polizzi S, Mahajan VB. Intravitreal anti-VEGF injections in pregnancy: Case series and review of literature. J Ocul Pharmacol Ther. 2015;31(10):605-610. PMID: 26302032.
- 4. Peracha ZH, Rosenfeld PJ. Anti-vascular endothelial growth factor therapy in pregnancy: What we know, what we don't know, and what we don't know we don't know. Retina. 2016;36(8):1413-1417. PMID: 27388726.
- 5. Sakai T, Mori C, Ohtsu F. 2022. Potential safety signal of pregnancy loss with vascular endothelial growth factor inhibitor intraocular injection: A disproportionality analysis using the Food and Drug Administration

  Adverse Event Reporting System. Front Pharmacol 13: 1063625. https://www.ncbi.nlm.nih.gov/pmc/articles/pmid/36438807/